#### AMENDMENTS TO THE CLAIMS

The Examiner is directed to the International Preliminary Examination Report (IPER) issued June 29, 2004 in which it is reported that Applicant's amendments to the claims filed on December 15, 2003 and April 28, 2004 under PCT Article 34 have been entered. For the convenience of the Examiner, copies of the December 15, 2003 and April 28, 2004 Amendments and the IPER are enclosed herewith, as Appendices B, C and D, respectively.

Currently amended claims 1, 37, 84 and 123 take into account amendments made in the Article 34 Amendments filed December 15, 2003 and April 28, 2004.

This listing of claims will replace all prior versions, and listings, of claims in the application:

## 1. (Currently Amended) A compound having the structure:

$$R_{11}$$

$$R_{10}$$

$$R_{11}$$

$$R_{11}$$

$$R_{10}$$

$$R_{11}$$

$$R_{2}$$

$$R_{3}$$

$$R_{4}$$

$$R_{7}$$

$$R_{6}$$

$$R_{7}$$

$$R_{6}$$

$$R_{11}$$

$$R_{3}$$

$$R_{2}$$

$$R_{4}$$

$$R_{5}$$

$$R_{7}$$

$$R_{6}$$

or pharmaceutically acceptable salt, ester, or salt of ester thereof;

wherein  $R_1$  is hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, or an aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl moiety; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms; or

Page 3 of 34

Express Mail No.: EV 314103555 US 3742671v1

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms;

R<sub>4</sub> is hydrogen or halogen;

R<sub>5</sub> is hydrogen, an oxygen protecting group or a prodrug moiety;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

n is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or an aliphatic moiety optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2-R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2-R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or - N(alkyl), or wherein  $X_2$ -R<sub>14</sub> together are N<sub>3</sub> or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is – (C=O)NHR<sub>15</sub> –(C=O)OR<sub>15</sub>, or –(C=O)R<sub>15</sub>, wherein each occurrence of R<sub>15</sub> is independently hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or R<sub>14</sub> is –SO<sub>2</sub>(R<sub>16</sub>), wherein R<sub>16</sub> is an aliphatic moiety, wherein one or more of R<sub>14</sub>, R<sub>15</sub>, or R<sub>16</sub> are optionally substituted with one or more occurrences of hydroxyl,

protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $\mathbf{R}_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;

 $\mathbf{R}_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH<sub>2</sub> or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and **Z** is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or aliphatic, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond;

with the proviso that when n is 1; X is O;  $R_1$  is methyl;  $R_2$ ,  $R_3$ ,  $R_7$  and  $R_{11}$  are each hydrogen;  $R_5$  is hydrogen,  $C_{1-4}$ alkyl or  $-C(=O)C_{1-4}$ alkyl;  $R_6$  is hydrogen, OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl; and  $R_9$  is OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl; then one or more if the following groups do not occur simultaneously as defined:

- (i) R<sub>4</sub> is hydrogen; R<sub>10</sub> and R<sub>8</sub> are independently OH, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1</sub>.

  4alkyl; and Y-Z is -CH<sub>2</sub>CH<sub>2</sub>- or -CH=CH-;
- (ii)  $R_4$  and  $R_8$  are each hydrogen;  $R_{10}$  is OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl; and Y-Z is  $-CHR^YCHR^Z$ -, -CH=CH- or ; wherein  $R^Y$  and  $R^Z$  are independently hydrogen,  $C_{1-4}$ alkyl or  $C_{1-4}$ alkanoyl; and
- (iii) R<sub>4</sub> and R<sub>10</sub> are each hydrogen, OH, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-4</sub>alkyl; R<sub>8</sub> is hydrogen, OH, halogen, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-4</sub>alkyl; and Y-Z is -CH<sub>2</sub>CH<sub>2</sub>-, -CH=CH- or -C(=O)CH<sub>2</sub>-.

2. (Original) The compound of claim 1, where the following groups do not occur simultaneously as defined:

X is oxygen,

R<sub>1</sub> is methyl,

R<sub>2</sub> and R<sub>3</sub> are each hydrogen,

R<sub>4</sub> is hydrogen,

 $R_5$  is hydrogen,  $C_{1-6}$ alkyl or  $C_{1-6}$ alkanoyl,

 $R_6$  is OR', where R' is hydrogen,  $C_{1-6}$ alkyl or  $C_{1-6}$ alkanoyl with S-configuration,

R<sub>7</sub> is hydrogen,

Y and Z together represent  $-CHR_{17}$ - $CHR_{18}$ -or  $-CR_{17}$ = $CR_{18}$ -, wherein  $R_{17}$  and  $R_{18}$  are independently hydrogen, or when Y and Z are  $-CHR_{17}$ - $CHR_{18}$ ,  $R_{17}$  and  $R_{18}$  taken together are -O-;

R<sub>8</sub> is hydrogen or OR', where R' is hydrogen, C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkanoyl,

R<sub>9</sub> is OR', where R' is hydrogen, C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkanoyl,

R<sub>10</sub> is OR", where R" is hydrogen, C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkanoyl; and

R<sup>11</sup> is hydrogen.

3. (Original) The compound of claim 1, wherein:

 $R_1$  is hydrogen, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched C<sub>1-6</sub>alkyl, straight or branched C<sub>1-6</sub>heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R<sub>4</sub> is hydrogen or halogen;

 $\mathbf{R}_5$  is hydrogen or a protecting group;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

**n** is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

R<sub>8</sub> is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or C<sub>1-6</sub>alkyl optionally substituted with hydroxyl, protected hydroxyl, SR<sub>12</sub>, or NR<sub>12</sub>R<sub>13</sub>;

R<sub>9</sub> is hydrogen, halogen, hydroxyl, protected hydroxyl, OR<sub>12</sub>, SR<sub>12</sub>, NR<sub>12</sub>R<sub>13</sub>, -X<sub>1</sub>(CH<sub>2</sub>)<sub>p</sub>X<sub>2</sub>-R<sub>14</sub>, or is C<sub>1-6</sub>alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or  $-X_1(CH_2)_pX_2-R_{14}$ ;

wherein R<sub>12</sub> and R<sub>13</sub> are, independently for each occurrence, hydrogen, C<sub>1-6</sub>alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or R<sub>12</sub> and R<sub>13</sub>, taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of R<sub>12</sub> and R<sub>13</sub> are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein X<sub>1</sub> and X<sub>2</sub> are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein X<sub>2</sub>-R<sub>14</sub> together are N<sub>3</sub> or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

R<sub>14</sub> is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is – (C=O)NHR<sub>15</sub> -(C=O)OR<sub>15</sub>, or -(C=O)R<sub>15</sub>, wherein each occurrence of  $R_{15}$  is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or R<sub>14</sub> is -SO<sub>2</sub>(R<sub>16</sub>), wherein R<sub>16</sub> is an alkyl moiety, wherein one or more of R<sub>14</sub>, R<sub>15</sub>, or R<sub>16</sub> are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $\mathbf{R}_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;

 $\mathbf{R}_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH<sub>2</sub> or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond.

- 4. (Original) The compound of claim 3, where X is oxygen and n is 1.
- 5. (Original) The compound of claim 3, where  $R_4$  is halogen.
- 6. (Original) The compound of claim 3, where R<sub>4</sub> is fluorine.
- 7. (Original) The compound of claim 3, where Y and Z together represent-CH=CH-
- 8. (Original) The compound of claim 3, where Y and Z together represent trans –CH=CH-.
- 9. (Original) The compound of claim 3, wherein  $R_1$  and  $R_2$  are each methyl and  $R_3$  is hydrogen and the compound has the structure:

Express Mail No.: EV 314103555 US 3742671v1

$$R_{11}$$
 $R_{10}$ 
 $R_{11}$ 
 $R_{10}$ 
 $R_{11}$ 
 $R$ 

wherein R<sub>4</sub>-R<sub>11,</sub> n, X, Y and Z are as defined in claim 3.

- 10. (Original) The compound of claim 9, wherein X is oxygen and n is 1.
- 11. (Original) The compound of claim 9, wherein R<sub>4</sub> is halogen.
- 12. (Original) The compound of claim 9, wherein Y and Z together represent -CH=CH.
- 13. **(Original)** The compound of claim 9, wherein X is oxygen, n is 1, R<sub>4</sub> is halogen and Y and Z together represent –CH=CH-.
- 14. (Original) The compound of claim 12 or 13 wherein –CH=CH- is trans.
- 15. (Original) The compound of claim 3, wherein  $R_9$  is  $NR_{12}R_{13}$  and the compound has the structure:

$$R_{12}$$
 $R_{13}$ 
 $R_{10}$ 
 $R_{10}$ 
 $R_{10}$ 
 $R_{11}$ 
 $R_{12}$ 
 $R_{13}$ 
 $R_{12}$ 
 $R_{13}$ 
 $R_{12}$ 
 $R_{13}$ 
 $R_{14}$ 
 $R_{15}$ 
 $R$ 

Page 9 of 34

Express Mail No.: EV 314103555 US 3742671v1

wherein R<sub>1</sub>-R<sub>12</sub>, n, X, Y and Z are as defined in claim 3, or

R<sub>13</sub> and R<sub>8</sub> may, when taken together, form a cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydrogen, alkyloxy, amino, alkylamino, aminoalkyl, and halogen.

- 16. (Original) The compound of claim 15, wherein X is oxygen and n is 1.
- 17. (Original) The compound of claim 15, wherein  $R_4$  is halogen.
- 18. (Original) The compound of claim 15, wherein Y and Z together represent –CH=CH-.
- 19. (Original) The compound of claim 15, wherein  $R_1$  and  $R_2$  are each methyl and  $R_3$  is hydrogen.
- 20. (Original) The compound of claim 15, wherein X is oxygen, n is 1,  $R_1$  and  $R_2$  are each methyl,  $R_3$  is hydrogen,  $R_4$  is halogen, and Y and Z together represent –CH=CH-.
- 21. (Original) The compound of claim 18 or 20, wherein –CH=CH- is trans.
- 22. (Currently Amended) A compound having the structure: The compound of claim 1 having the structure:

Page 10 of 34

Express Mail No.: EV 314103555 US
Attorney Docket No.: 2003946-0110
3742671v1
Client Reference: ANDI/US

Page 11 of 34

Express Mail No.: EV 314103555 US 3742671v1

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

### Claims 23-36 (Canceled)

# 37. (Currently Amended) A pharmaceutical composition comprising: a compound having the structure: of any one of claims 1, 9 and 15;

$$\begin{array}{c|c}
R_{10} & O & R_1 \\
R_{11} & X & R_2 \\
R_{11} & X & R_4 \\
\hline
R_{11} & X & R_6 \\
\hline
R_{12} & X & R_6 \\
\hline
R_{13} & X & R_6 \\
\hline
R_{14} & X & R_6 \\
\hline
R_{15} & X & R_6 \\
\hline
R_{1$$

or pharmaceutically acceptable salt, ester, or salt of ester thereof;

wherein R<sub>1</sub> is hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl;

R<sub>2</sub>-and-R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, or an aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl moiety; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms; or

R<sub>1</sub>-and R<sub>3</sub>, when taken-together, may-form-a substituted or unsubstituted; saturated or unsubstituted or unsubstituted.

R<sub>4</sub> is hydrogen or halogen;

Rs is hydrogen, an oxygen protecting group or a prodrug moiety;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

n is 0 2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

R<sub>8</sub> is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or an aliphatic moiety optionally substituted with hydroxyl, protected hydroxyl, SR<sub>12</sub>, or NR<sub>12</sub>R<sub>13</sub>;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ ,  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or  $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein R<sub>12</sub> and R<sub>13</sub> are, independently for each occurrence, hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or a protecting group, or R<sub>12</sub> and R<sub>13</sub>, taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of R<sub>12</sub> and R<sub>13</sub> are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein X<sub>1</sub> and X<sub>2</sub> are each independently absent, or are oxygen, NH, or N(alkyl), or wherein X<sub>2</sub>-R<sub>14</sub>-together are N<sub>3</sub> or are a saturated or unsaturated heterocyclic moiety,

p is 2 10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $(C=O)NHR_{15}$ — $(C=O)OR_{15}$ , or  $(C=O)R_{15}$ , wherein each occurrence of  $R_{15}$ —is independently hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or  $R_{14}$  is  $SO_2(R_{16})$ , wherein  $R_{16}$  is an aliphatic moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

R<sub>10</sub> is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;

**R**<sub>11</sub> is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N alkyl, CH2 or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or aliphatic, or R<sub>17</sub> and R<sub>18</sub> taken together is O, CH<sub>2</sub> or NR<sub>19</sub>, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and

a pharmaceutically acceptable earrier; carrier.

with the proviso that when n is 1; X is O; R<sub>1</sub> is methyl; R<sub>2</sub>, R<sub>3</sub>, R<sub>7</sub> and R<sub>11</sub> are each hydrogen; R<sub>5</sub> is hydrogen, C<sub>1.4</sub>alkyl or C(=O)C<sub>1.4</sub>alkyl; R<sub>6</sub> is hydrogen, OH, C<sub>1.4</sub>alkoxy or OC(=O)C<sub>1.4</sub>alkyl; and R<sub>9</sub> is OH, C<sub>1.4</sub>alkoxy or OC(=O)C<sub>1.4</sub>alkyl; then one or more if the following groups do not occur simultaneously as defined:

- (i) R<sub>4</sub> is hydrogen; R<sub>10</sub> and R<sub>8</sub> are independently OH, C<sub>1-4</sub>alkoxy or OC(=O)C<sub>1-4</sub>alkyl; and Y Z is CH<sub>2</sub>CH<sub>2</sub> or CH=CH;
- (ii) R4 and R8 are each hydrogen; R10 is OH, C1 4alkoxy or OC(=0)C1 4alkyl; and Y

- (iii)  $R_4$  and  $R_{10}$  are each hydrogen, OH,  $C_{1-4}$ alkoxy or  $OC(=O)C_{1-4}$ alkyl;  $R_8$  is hydrogen, OH, halogen,  $C_{1-4}$ alkoxy or  $OC(=O)C_{1-4}$ alkyl; and Y Z is  $CH_2CH_2$ , CH=CH or  $C(=O)CH_2$ .
- 38. (Original) The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to inhibit NF-κB activation.
- 39. (Original) The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to inhibit AP-1 activation.
- 40. (**Original**) The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to inhibit a protein kinase.

Page 14 of 34

Express Mail No.: EV 314103555 US

Attorney Docket No.: 2003946-0110
3742671v1

Client Reference: ANDI/US

41. (Currently Amended) The pharmaceutical composition of elaim 39 claim 40, wherein

the protein kinase is MEKK1, MEK1, VEGFr or PDGFr.

42. (Original) The pharmaceutical composition of claim 37, wherein the compound is

present in an amount effective to inhibit proliferation of cancerous cells and angiogenesis on

solid tumors.

43. (Original) The pharmaceutical composition of claim 37, wherein the compound is

present in an amount effective to have an anti-inflammatory effect.

44. (Original) The pharmaceutical composition of claim 37, wherein the compound is

present in an amount effective to treat psoriasis.

45. (Original) The pharmaceutical composition of claim 37, wherein the compound is

present in an amount effective to reduce skin photodamage.

46. (Original) The pharmaceutical composition of claim 37, wherein the compound is

present in an amount effective to prevent restenosis.

Claims 47-65 (Canceled)

66. (Currently Amended) A pharmaceutical composition comprising: The pharmaceutical

composition of claim 37 wherein the compound has the structure:

a compound having the structure:

Page 16 of 34

Express Mail No.: EV 314103555 US 3742671v1

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable earrier.

## Claims 67-80 (Canceled)

81. (Currently Amended) A topical pharmaceutical composition for preventing or treating UVB-induced photodamage comprising:

a compound having the structure:

$$R_{11}$$

$$R_{10}$$

$$R_{11}$$

$$R$$

or pharmaceutically acceptable salt, ester, or salt of ester thereof;

wherein  $R_1$  is hydrogen, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ 6heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

 $R_2$  and  $R_3$  are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R<sub>4</sub> is hydrogen or halogen;

R<sub>5</sub> is hydrogen or a protecting group hydrogen, an oxygen protecting group or a prodrug moiety;

 $\mathbf{R}_{6}$  is hydrogen, hydroxyl, or protected hydroxyl;

**n** is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2-R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2-R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1-6}$ alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or - N(alkyl), or wherein  $X_2$ -R<sub>14</sub> together are N<sub>3</sub> or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is – (C=O)NHR<sub>15</sub> –(C=O)OR<sub>15</sub>, or –(C=O)R<sub>15</sub>, wherein each occurrence of R<sub>15</sub> is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or R<sub>14</sub> is –SO<sub>2</sub>(R<sub>16</sub>), wherein R<sub>16</sub> is an alkyl moiety, wherein one or more of R<sub>14</sub>, R<sub>15</sub>, or

R<sub>16</sub> are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $\mathbf{R}_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;

 $\mathbf{R}_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH<sub>2</sub> or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and

a pharmaceutically acceptable carrier;

wherein the compound is present in an amount effective to prevent or treat UVB-induced photodamage.

- 82. (Original) The pharmaceutical composition of claim 81, further comprising a cosmetic ingredient.
- 83. (Original) The pharmaceutical composition of claim 82, wherein the cosmetic ingredient is a sunscreen.
- 84. (Currently Amended) A method for treating an inflammatory and/or autoimmune disorder or a disorder resulting from increased angiogenesis and/or cell proliferation comprising: administering to a subject in need thereof a therapeutically effective amount of a compound having the structure: of any one of claims 1, 9 and 15;

Page 19 of 34

$$R_{11}$$
 $R_{10}$ 
 $R_{10}$ 
 $R_{10}$ 
 $R_{11}$ 
 $R_{2}$ 
 $R_{3}$ 
 $R_{2}$ 
 $R_{4}$ 
 $R_{8}$ 
 $R_{7}$ 
 $R_{8}$ 
 $R_{8}$ 
 $R_{11}$ 
 $R_{10}$ 
 $R_{11}$ 
 $R_{11}$ 
 $R_{11}$ 
 $R_{11}$ 
 $R_{12}$ 
 $R_{12}$ 
 $R_{13}$ 
 $R_{14}$ 
 $R_{15}$ 
 $R_{15}$ 

or pharmaceutically acceptable salt, ester, or salt of ester thereof;

wherein  $R_1$  is hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, or an aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl moiety; or

R<sub>1</sub>-and R<sub>2</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms; or

R<sub>1</sub>-and R<sub>3</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsubstituted or unsubstituted, saturated or unsubstitute

R4 is hydrogen or halogen;

R<sub>5</sub> is hydrogen, an oxygen protecting group or a prodrug;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

n is 0.2;

R<sub>2</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

R<sub>8</sub> is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or an aliphatic moiety optionally substituted with hydroxyl, protected hydroxyl, SR<sub>12</sub>, or NR<sub>12</sub>R<sub>13</sub>;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ ,  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or  $X_1(CH_2)_pX_2$ - $R_{14}$ ;

Express Mail No.: EV 314103555 US
Attorney Docket No.: 2003946-0110
3742671v1
Client Reference: ANDI/US

wherein R<sub>12</sub> and R<sub>13</sub> are, independently for each occurrence, hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or a protecting group, or R<sub>12</sub> and R<sub>13</sub>, taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of R<sub>12</sub> and R<sub>13</sub> are optionally further—substituted—with—one—or—more—occurrences—of—hydroxyl, protected—hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or N(alkyl), or wherein  $X_2$   $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

<del>p is 2-10, and</del>

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $(C=O)NHR_{15}$   $(C=O)OR_{15}$ , or  $(C=O)R_{15}$ , wherein each occurrence of  $R_{15}$  is independently hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or  $R_{14}$  is  $SO_2(R_{16})$ , wherein  $R_{16}$  is an aliphatic moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

R<sub>10</sub> is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;

**R**<sub>11</sub>-is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;

¥ is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or aliphatic, or R<sub>17</sub> and R<sub>18</sub> taken together is O, CH<sub>2</sub> or NR<sub>19</sub>, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and

a pharmaceutically acceptable carrier or diluent;

with the proviso that when n is 1; X is O; R<sub>1</sub> is methyl; R<sub>2</sub>, R<sub>3</sub>, R<sub>2</sub> and R<sub>11</sub> are each hydrogen; R<sub>5</sub> is hydrogen, C<sub>1-4</sub>alkyl or C(=O)C<sub>1-4</sub>alkyl; R<sub>6</sub> is hydrogen, OH, C<sub>1-4</sub>alkoxy or OC(=O)C<sub>1-4</sub>alkyl; and R<sub>9</sub> is OH, C<sub>1-4</sub>alkoxy or OC(=O)C<sub>1-4</sub>alkyl; then one or more if the following groups do not occur simultaneously as defined:

- (i) R<sub>4</sub> is hydrogen; R<sub>10</sub> and R<sub>8</sub> are independently OH, C<sub>1</sub> 4alkoxy or OC(=O)C<sub>1</sub>.

  4alkyl; and Y-Z is -CH<sub>2</sub>CH<sub>2</sub> or -CH=CH; and
- Z is CHR<sup>4</sup>CHR<sup>2</sup>, CH=CH or ; wherein R<sup>4</sup> and R<sup>2</sup> are independently hydrogen, C<sub>1</sub> 4alkyl or C<sub>1</sub> 4alkanoyl; and
- (iii) R<sub>4</sub>-and R<sub>10</sub>-are each hydrogen, OH, C<sub>1</sub>-alkoxy or OC(=O)C<sub>1</sub>-alkyl; R<sub>8</sub> is hydrogen, OH, halogen, C<sub>1</sub>-alkoxy or OC(=O)C<sub>1</sub>-alkyl; and Y Z is CH<sub>2</sub>CH<sub>2</sub>-, CH=CH-or C(=O)CH<sub>2</sub>-; whereby the compound induces mRNA degradation and the method is for treating a disorder resulting from cell proliferation.
- 85. (Original) The method of claim 84, wherein the method is for treating a disorder selected from the group consisting of rheumatoid arthritis, psoriasis, asthma, cancer, sepsis, inflammatory bowel disease, atopic dermatitis, Crohn's disease, and autoimmune disorders.
- 86. (Original) The method of claim 84, wherein the method is for treating rheumatoid arthritis.
- 87. (Original) The method of claim 84, wherein the method is for treating psoriasis.
- 88. (Original) The method of claim 84, wherein the method is for treating asthma.

Claims 89-107 (Canceled)

# 108. (Currently Amended) The method of claim 84, comprising administering a compound having wherein the compound has the structure:

Express Mail No.: EV 314103555 US 3742671v1

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

## Claims 109-118 (Canceled)

119. (Currently Amended) A method for providing protection against UVB-induced photodamage to a subject, said method comprising:

Administering to the subject in need thereof a composition comprising a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof;

wherein  $R_1$  is hydrogen, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ 6heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

 $R_2$  and  $R_3$  are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R<sub>4</sub> is hydrogen or halogen;

R<sub>5</sub> is hydrogen or a protecting group hydrogen, an oxygen protecting group or a prodrug moiety;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

**n** is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $\mathbf{R_9}$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2-R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2-R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1-6}$ alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or - N(alkyl), or wherein  $X_2$ -R<sub>14</sub> together are N<sub>3</sub> or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is – (C=O)NHR<sub>15</sub> –(C=O)OR<sub>15</sub>, or –(C=O)R<sub>15</sub>, wherein each occurrence of R<sub>15</sub> is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or R<sub>14</sub> is –SO<sub>2</sub>(R<sub>16</sub>), wherein R<sub>16</sub> is an alkyl moiety, wherein one or more of R<sub>14</sub>, R<sub>15</sub>, or R<sub>16</sub> are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

R<sub>10</sub> is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;

 $\mathbf{R}_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH<sub>2</sub> or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and **Z** is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and

a pharmaceutically acceptable carrier or diluent.

- 120. **(Original)** The method of claim 119, wherein in the step of administering, the composition is administered topically.
- 121. (Original) The method of claim 119, wherein the photodamage is skin wrinkles.
- 122. (Original) The method of claim 119, wherein the photodamage is a skin cancer.
- 123. (Currently Amended) A method for preventing or reducing the rate of restenosis, comprising:

inserting a stent into an obstructed blood vessel, the stent having a generally tubular structure, the surface of the structure being coated with (or otherwise adapted to release) a composition comprising a compound having the structure:

$$R_{11}$$

$$R_{9}$$

$$R_{10}$$

$$R_{10}$$

$$R_{11}$$

$$R_$$

or pharmaceutically acceptable salt, ester, or salt of ester thereof;

wherein  $R_1$  is hydrogen, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ 6heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

 $R_2$  and  $R_3$  are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R<sub>4</sub> is hydrogen or halogen;

R<sub>5</sub> is hydrogen or a protecting group hydrogen, an oxygen protecting group or a prodrug moiety;

**R**<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl; **n** is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2-R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2-R_{14}$ ;

wherein R<sub>12</sub> and R<sub>13</sub> are, independently for each occurrence, hydrogen, C<sub>1-6</sub>alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or R<sub>12</sub> and R<sub>13</sub>, taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of R<sub>12</sub> and R<sub>13</sub> are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or - N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is – (C=O)NHR<sub>15</sub> –(C=O)OR<sub>15</sub>, or –(C=O)R<sub>15</sub>, wherein each occurrence of R<sub>15</sub> is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or R<sub>14</sub> is –SO<sub>2</sub>(R<sub>16</sub>), wherein R<sub>16</sub> is an alkyl moiety, wherein one or more of R<sub>14</sub>, R<sub>15</sub>, or R<sub>16</sub> are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

R<sub>10</sub> is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;

 $\mathbf{R}_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH<sub>2</sub> or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and **Z** is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and optionally

a pharmaceutically acceptable carrier or diluent;

such that the obstruction is eliminated and the composition is delivered in amounts effective to prevent or reduce the rate of restenosis;

with the proviso that the following groups do not occur simultaneously as defined: n is 1; X is O;  $R_1$  is methyl;  $R_2$ ,  $R_3$ ,  $R_4$ ,  $R_7$ ,  $R_8$  and  $R_{11}$  are each hydrogen;  $R_5$  is hydrogen,  $C_{1-4}$ alkyl or  $-C(=O)C_{1-4}$ alkyl;  $R_6$  is hydrogen, OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl;  $R_9$  and  $R_{10}$  are independently OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl; and Y-Z is  $-CHR^YCHR^Z$ -, -CH=CH- or

wherein 
$$R^{Y}$$
 and  $R^{Z}$  are independently hydrogen,  $C_{1-4}$ alkyl or  $C_{1-4}$ alkanoyl.

124. (Currently Amended) A method for expanding the lumen of a body passageway, comprising:

inserting a stent into the passageway, the stent having a generally tubular structure, the surface of the structure being coated with (or otherwise adapted to release) a composition comprising a compound having the structure:

$$R_{11}$$

$$R_{10}$$

$$R_{11}$$

$$R_{10}$$

$$R_{11}$$

$$R_{2}$$

$$R_{3}$$

$$R_{2}$$

$$R_{4}$$

$$R_{5}$$

$$R_{7}$$

$$R_{6}$$

$$R_{6}$$

$$R_{10}$$

$$R_{11}$$

$$R_{2}$$

$$R_{2}$$

$$R_{3}$$

$$R_{4}$$

$$R_{5}$$

$$R_{7}$$

$$R_{6}$$

$$R_{10}$$

or pharmaceutically acceptable salt, ester, or salt of ester thereof;

wherein  $R_1$  is hydrogen, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched C<sub>1-6</sub>alkyl, straight or branched C<sub>1-6</sub>heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R<sub>4</sub> is hydrogen or halogen;

 $\mathbf{R}_5$  is hydrogen or a protecting group;

 $\mathbf{R}_{6}$  is hydrogen, hydroxyl, or protected hydroxyl;

n is 0-2;

 $\mathbf{R}_7$ , for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $\mathbf{R}_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2-R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2-R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1-6}$ alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or - N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is – (C=O)NHR<sub>15</sub> –(C=O)OR<sub>15</sub>, or –(C=O)R<sub>15</sub>, wherein each occurrence of R<sub>15</sub> is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or R<sub>14</sub> is –SO<sub>2</sub>(R<sub>16</sub>), wherein R<sub>16</sub> is an alkyl moiety, wherein one or more of R<sub>14</sub>, R<sub>15</sub>, or R<sub>16</sub> are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $\mathbf{R}_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;

 $\mathbf{R}_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH<sub>2</sub> or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of  $R_{17}$  and  $R_{18}$  is independently hydrogen or  $C_{1.6}$ alkyl, or  $R_{17}$  and  $R_{18}$  taken

together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein  $R_{19}$  is hydrogen or  $C_{1-6}$ alkyl, and Y and Z may be connected by a single or double bond; and optionally

a pharmaceutically acceptable carrier or diluent; such that the passageway is expanded.

- 125. (Original) The method of claim 124, wherein the lumen of a body passageway is expanded in order to eliminate a biliary, gastrointestinal, esophageal, tracheal/bronchial, urethral and/or vascular obstruction.
- 126. (Original) The method of claim 125, wherein the lumen of a body passageway is expanded in order to eliminate a vascular obstruction.

Page 32 of 34

Express Mail No.: EV 314103555 US
Attorney Docket No.: 2003946-0110
3742671v1
Client Reference: ANDI/US